Rooke 10/626,571

05/10/2004

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L9 ANSWER 1 OF 7 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER:

2004:397248 HCAPLUS

TITLE:

Reduced calcification of bioprostheses, cross-linked

via an improved carbodiimide based method

AUTHOR (S):

Everaerts, Frank; Torrianni, Mark;

van Luyn, Marja; van Wachem, Pauline; Feijen, Jan;

Hendriks, Mark

CORPORATE SOURCE:

Biomaterials S&T, Medtronic Bakken Research Center,

Maastricht, 6229 GW, Neth.

SOURCE:

Biomaterials (2004), 25(24), 5523-5530

CODEN: BIMADU; ISSN: 0142-9612

PUBLISHER:

Elsevier Science Ltd.

DOCUMENT TYPE:

Journal English

LANGUAGE:

Glutaraldehyde fixation of bioprosthetic tissue has been used successfully for almost 40 yr. However, it is generally recognized that glutaraldehyde fixation of bioprostheses is associated with the occurrence of calcification. Accordingly, many efforts have been undertaken to develop techniques for the fixation of bioprostheses, which will not lead to calcification. Here we describe a new improved carbodiimide based crosslinking method. Rather than crosslinking the tissue through its free primary amine groups, these groups were first blocked with butanal and the tissue was then cross-linked by means of carbodiimide activation of tissue carboxylic acid groups followed by a reaction with a poly(propylene glycol)bis 2-(aminopropyl) ether, (Jeffamine). It was demonstrated that cross-linked porcine leaflets had a calcification of less than 1 mg/g

CC 63-7 (Pharmaceuticals)

calcification model used.

- ST carbodiimide heart valve bioprosthetic
- IT Calcification

## Crosslinking

(reduced calcification of bioprostheses, cross-linked via an improved carbodiimide based method)

tissue after 8 wk sub-dermal implantion in rats. Furthermore, aortic wall calcification was reduced to 50 mg/g, compared to standard glutaraldehyde fixed tissue, which showed 120 mg/g tissue calcification in the 8 wk

IT Heart

(valve, bioprosthetic; reduced calcification of bioprostheses, cross-linked via an improved carbodiimide based method)

IT 7440-70-2, Calcium 7723-14-0, Phosphorus

RL: BSU (Biological study, unclassified); BIOL (Biological study) (reduced calcification of bioprostheses, cross-linked via an improved carbodiimide based method)

IT 123-72-8, Butanal

RL: PEP (Physical, engineering or chemical process); PRP (Properties); PYP (Physical process); PROC (Process)

(reduced calcification of bioprostheses, cross-linked via an improved carbodiimide based method)

IT 9046-10-0, Jeffamine 400

RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(reduced calcification of bioprostheses, cross-linked via an improved carbodiimide based method)

IT 1892-57-5, 1-Ethyl-3-(3-dimethylaminopropyl)carbodiimide 6066-82-6, N-Hydroxysuccinimide

RL: RCT (Reactant); RACT (Reactant or reagent)

(reduced calcification of bioprostheses, cross-linked via an improved carbodiimide based method)

REFERENCE COUNT: 20 THERE ARE 20 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 2 OF 7 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 2003:950040 HCAPLUS

DOCUMENT NUMBER: 140:19764

TITLE: Methods of inducing the expression of bone

morphogenetic proteins (BMPs) and transforming growth

factor-beta proteins (TGF- $\beta$ s) in cells

INVENTOR(S): Mckay, William F.; Boden, Scott D.; Yoon,

Sangwook T.

PATENT ASSIGNEE(S): USA

SOURCE: U.S. Pat. Appl. Publ., 81 pp., Cont.-in-part of U.S.

Ser. No. 292,951.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT NO.	KIND	DATE	APPLICATION NO.		DATE		
US 2003225021	A1	20031204	US 2003-382844		20030307		
US 2003180266	<b>A1</b>	20030925	US 2002-292951		20021113		
PRIORITY APPLN. INFO.:		•	US 2001-331321P	P	20011114		
			US 2002-292951	A2	20021113		
			US 1988-124238	Α	19880729		
			US 2000-959578	Α	20000428		

- AB A method of inducing the expression of one or more bone morphogenetic proteins and/or transforming growth factor-β proteins in a cell is described. The method includes transfecting a cell with an isolated nucleic acid comprising a nucleotide sequence encoding a LIM mineralization protein operably linked to a promoter. The one or more bone morphogenetic proteins can be BMP-2, BMP-4, BMP-6, BMP-7 or combinations thereof. The transforming growth factor-β protein can be transforming growth factor-β1 protein (TGF-β1). Transfection may be accomplished ex vivo or in vivo by direct injection of virus or naked DNA, or by a nonviral vector such as a plasmid. The method can be used to induce bone formation in osseous cells or to stimulate proteoglycan and/or collagen production in cells capable of producing proteoglycyan and/or collagen (e.g., intervertebral disk cells).
- IC ICM A61K048-00
  - ICS C12N005-08; C12N015-861; C12N015-867
- NCL 514044000; 424093200; 435456000; 435366000
- CC 63-1 (Pharmaceuticals)
  - Section cross-reference(s): 3, 6, 14
- ST bone morphogenetic protein transforming growth factor gene therapy; BMP TGF gene bone formation intervertebral disk disease; oligonucleotide transformation cell implant bone disease
- IT Bone morphogenetic proteins
  - RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
    - (2; methods of inducing the expression of **bone** morphogenetic proteins (BMPs) and transforming growth factor-beta proteins (TGF- $\beta$ s) in cells)
- IT Bone morphogenetic proteins
  - RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

```
(4; methods of inducing the expression of bone morphogenetic
        proteins (BMPs) and transforming growth factor-beta proteins
        (TGF-βs) in cells)
     Bone morphogenetic proteins
IT
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (6; methods of inducing the expression of bone morphogenetic
        proteins (BMPs) and transforming growth factor-beta proteins
        (TGF-\beta s) in cells)
     Bone morphogenetic proteins
IT
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (7; methods of inducing the expression of bone morphogenetic
        proteins (BMPs) and transforming growth factor-beta proteins
        (TGF-βs) in cells)
IT
     Adenoviridae
        (AdLMP-1; methods of inducing the expression of bone
        morphogenetic proteins (BMPs) and transforming growth factor-beta
        proteins (TGF-\betas) in cells)
     Gene, animal
IT
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (BMP-2; methods of inducing the expression of bone
        morphogenetic proteins (BMPs) and transforming growth factor-beta
        proteins (TGF-βs) in cells)
IT
     Gene, animal
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (BMP-4; methods of inducing the expression of bone
        morphogenetic proteins (BMPs) and transforming growth factor-beta
        proteins (TGF-\betas) in cells)
     Gene, animal
IT
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (BMP-6; methods of inducing the expression of bone
        morphogenetic proteins (BMPs) and transforming growth factor-beta
       proteins (TGF-\betas) in cells)
IT
     Gene, animal
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (BMP-7; methods of inducing the expression of bone
        morphogenetic proteins (BMPs) and transforming growth factor-beta
       proteins (TGF-\betas) in cells)
IT
     Proteins
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (HLMP-1; methods of inducing the expression of bone
       morphogenetic proteins (BMPs) and transforming growth factor-beta
       proteins (TGF-\betas) in cells)
ΙT
     Proteins
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (HLMP-1s; methods of inducing the expression of bone
       morphogenetic proteins (BMPs) and transforming growth factor-beta
       proteins (TGF-\betas) in cells)
IT
     Proteins
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (HLMP-2; methods of inducing the expression of bone
       morphogenetic proteins (BMPs) and transforming growth factor-beta
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proteins (TGF-βs) in cells)
     Proteins
IT
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (HLMP-3; methods of inducing the expression of bone
        morphogenetic proteins (BMPs) and transforming growth factor-beta
        proteins (TGF-βs) in cells)
     Proteins
IT
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (LIM domain-containing; methods of inducing the expression of bone
        morphogenetic proteins (BMPs) and transforming growth factor-beta
        proteins (TGF-βs) in cells)
IT
     Proteins
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (LMP-1; methods of inducing the expression of bone
        morphogenetic proteins (BMPs) and transforming growth factor-beta
        proteins (TGF-\betas) in cells)
IT
     Proteins
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (RLMP; methods of inducing the expression of bone
        morphogenetic proteins (BMPs) and transforming growth factor-beta
        proteins (TGF-βs) in cells)
IT
     Gene, animal
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (TGF-β1; methods of inducing the expression of bone
        morphogenetic proteins (BMPs) and transforming growth factor-beta
        proteins (TGF-\betas) in cells)
IT
     Cell
        (annulus fibrosus; methods of inducing the expression of bone
        morphogenetic proteins (BMPs) and transforming growth factor-beta
        proteins (TGF-\betas) in cells)
     Spinal column, disease
IT
        (intervertebral disk hernia; methods of inducing the expression of
        bone morphogenetic proteins (BMPs) and transforming growth
        factor-beta proteins (TGF-βs) in cells)
IT
     Spinal column
        (intervertebral disk; methods of inducing the expression of
        bone morphogenetic proteins (BMPs) and transforming growth
        factor-beta proteins (TGF-\betas) in cells)
IT
     Stem cell
        (mesenchymal; methods of inducing the expression of bone
        morphogenetic proteins (BMPs) and transforming growth factor-beta
        proteins (TGF-\betas) in cells)
     Adenoviral vectors
IT
       Bone formation
     Gene therapy
     Genetic vectors
     Hematopoietic precursor cell
     Mammalia
     Mesenchyme
     Molecular cloning
     Nucleic acid hybridization
     Ore genesis
     Plasmid vectors
     Retroviral vectors
     Retroviridae
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Transformation, genetic
     Transplant and Transplantation
     Viral vectors
     Virus
        (methods of inducing the expression of bone morphogenetic
        proteins (BMPs) and transforming growth factor-beta proteins
        (TGF-\beta s) in cells)
     Oligonucleotides
ΙT
     RL: BSU (Biological study, unclassified); PRP (Properties); BIOL
     (Biological study)
        (methods of inducing the expression of bone morphogenetic
        proteins (BMPs) and transforming growth factor-beta proteins
        (TGF-\beta s) in cells)
     Bone morphogenetic proteins
TΤ
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (methods of inducing the expression of bone morphogenetic
        proteins (BMPs) and transforming growth factor-beta proteins
        (TGF-βs) in cells)
     Nucleic acids
IT
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (methods of inducing the expression of bone morphogenetic
        proteins (BMPs) and transforming growth factor-beta proteins
        (TGF-\betas) in cells)
     Promoter (genetic element)
TΤ
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (methods of inducing the expression of bone morphogenetic
        proteins (BMPs) and transforming growth factor-beta proteins
        (TGF-\beta s) in cells)
TT
     Cytomegalovirus
        (promoter; methods of inducing the expression of bone
        morphogenetic proteins (BMPs) and transforming growth factor-beta
        proteins (TGF-\betas) in cells)
IT
     Cell nucleus
        (pulposus; methods of inducing the expression of bone
        morphogenetic proteins (BMPs) and transforming growth factor-beta
        proteins (TGF-\betas) in cells)
     Animal cell
IT
        (somatic; methods of inducing the expression of bone
        morphogenetic proteins (BMPs) and transforming growth factor-beta
        proteins (TGF-βs) in cells)
IT
     Transforming growth factors
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (\beta-; methods of inducing the expression of bone
        morphogenetic proteins (BMPs) and transforming growth factor-beta
        proteins (TGF-βs) in cells)
IT
     630143-01-0
                   630143-02-1
     RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (oligonucleotide; methods of inducing the expression of bone
        morphogenetic proteins (BMPs) and transforming growth factor-beta
        proteins (TGF-βs) in cells)
                                  630150-78-6
IT
     630150-76-4
                   630150-77-5
                                                630150-79-7
                                                              630150-80-0
     630150-81-1
                   630150-82-2
                                  630150-83-3
                                                630150-85-5
                                                              630150-86-6
     630150-87-7
                   63015.0-88-8
                                  630150-89-9
                                                630150-90-2
                                                              630150-91-3
     630150-92-4
                   630150-93-5
                                  630150-94-6
                                                630150-95-7
                                                              630150-96-8
     630150-97-9
                   630150-98-0
                                 630150-99-1
                                                630151-00-7
                                                              630151-01-8
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630151-04-1
                                               630151-06-3
                                                             630151-07-4
     630151-02-9
                   630151-03-0
                                 630151-12-1 630151-13-2
     630151-08-5
                   630151-10-9
     RL: PRP (Properties)
        (unclaimed nucleotide sequence; methods of inducing the expression of
        bone morphogenetic proteins (BMPs) and transforming growth
        factor-beta proteins (TGF-βs) in cells)
                                                             630151-11-0
                                630151-05-2
                                              630151-09-6
IT
     630150-75-3
                   630150-84-4
     RL: PRP (Properties)
        (unclaimed protein sequence; methods of inducing the expression of
        bone morphogenetic proteins (BMPs) and transforming growth
        factor-beta proteins (TGF-βs) in cells)
                                               630151-17-6 630151-18-7
IT
     630151-14-3
                 630151-15-4 630151-16-5
     630151-19-8
                   630151-20-1 630151-21-2
                                               630151-22-3
     RL: PRP (Properties)
        (unclaimed sequence; methods of inducing the expression of bone
        morphogenetic proteins (BMPs) and transforming growth factor-beta
       proteins (TGF-βs) in cells)
     ANSWER 3 OF 7 HCAPLUS COPYRIGHT 2004 ACS on STN
                        2003:397004 HCAPLUS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                         138:397329
                         cDNAs encoding rat and human LIM mineralization
TITLE:
                         proteins and their use in treatment of disk
                         degeneration and disk injury
                         McKay, William F.; Boden, Scott D.; Yoon,
INVENTOR (S):
                         Sangwook T.
                         Medtronic Sofamor Danek, USA
PATENT ASSIGNEE(S):
                         PCT Int. Appl., 94 pp.
SOURCE: ·
                         CODEN: PIXXD2
DOCUMENT TYPE:
                         Patent
                         English
LANGUAGE:
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
                       KIND
                                DATE
                                          APPLICATION NO.
                                                                  DATE
     PATENT NO.
                                -----
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     _____
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     WO 2003042368
                         A2
                                          WO 2002-US36465
                                                                  20021114
                                20030522
                                20031016
     WO 2003042368
                         A3
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
             CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
             LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
             PL, PT, RO, RU, SC, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT,
             TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ,
             MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG,
             CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
             PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR,
             NE, SN, TD, TG
                                            US 2002-292951
     US 2003180266
                         A1
                                20030925
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US 2000-959578 A 20000428

AB Methods of expressing LIM mineralization protein in non-osseous mammalian cells, such as stem cells or intervertebral disk cells (e.g., cells of the annulus fibrosus, or cells of the nucleus pulposus) are described. The methods involve transfecting the cells with an isolated nucleic acid comprising a nucleotide sequence encoding a LIM mineralization protein operably linked to a promoter. Transfection may be accomplished ex vivo

PRIORITY APPLN. INFO.:

US 2001-331321P

US 2002-292951

US 1988-124238

P 20011114

A 20021113

A 19880729

or in vivo by direct injection of virus or naked DNA, or by a nonviral vector such as a plasmid. Expression of the LIM mineralization protein can stimulate proteoglycan and/or collagen production in cells capable of producing proteoglycan and/or collagen. Methods for treating disk disease associated with trauma or disk degeneration are also described.

- IC ICM C12N
- CC 3-3 (Biochemical Genetics)

Section cross-reference(s): 1, 6, 13

- ST cDNA LIM mineralization protein human rat sequence; disk degeneration injury therapy LMP protein splicing isoform
- IT Bone morphogenetic proteins
  - RL: BSU (Biological study, unclassified); BIOL (Biological study)
    (2, LMP protein in stimulating synthesis of; cDNAs encoding rat and
    human LIM mineralization proteins and their use in treatment of disk
    degeneration and disk injury)
- IT Adenoviral vectors

(AdHLMP-1, LIM mineralization protein cDNA cloning in; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Protein motifs

(LIM domain, in LMP proteins; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Plasmid vectors

Retroviral vectors

Viral vectors

(LIM mineralization protein cDNA cloning in; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Proteoglycans, biological studies

RL: BSU (Biological study, unclassified); BIOL (Biological study)
(LIM mineralization protein in stimulating synthesis of; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT mRNA

RL: ANT (Analyte); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study)

(LIM mineralization protein; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT cDNA

RL: BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (LIM mineralization protein; cDNAs encoding rat and human LIM

mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Proteins

RL: BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(LMP (LIM mineralization protein), isoforms 1,2 and 3, of rat and human; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Bone morphogenetic proteins

RL: BSU (Biological study, unclassified); BIOL (Biological study)
(LMP protein in inducing synthesis of; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Osteocalcins

RL: BSU (Biological study, unclassified); BIOL (Biological study)

(LMP protein in stimulating secretion of; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Collagens, biological studies

RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(LMP protein in stimulating synthesis of, as carrier for LMP protein implant in vertebral disk; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Aggrecans

RL: BSU (Biological study, unclassified); BIOL (Biological study)
(LMP protein in stimulating synthesis of; cDNAs encoding rat and human
LIM mineralization proteins and their use in treatment of disk
degeneration and disk injury)

IT Bone formation

(LMP protein in; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT RNA splicing

(LMP protein mRNA; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Body, anatomical

(back, disease, pain, lower; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Pain

(back, lower; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Animal cell

Gene therapy

Human

Mammalia

Nucleic acid hybridization

Rattus

(cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Promoter (genetic element)

RL: BSU (Biological study, unclassified); BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(cytomegalovirus, for LMP proteins synthesis; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Bone, disease

(degenerative disk disease, spine stenosis; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Probes (nucleic acid)

RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); ANST (Analytical study); BIOL (Biological study); USES (Uses)

(for LMP protein cDNA; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Primers (nucleic acid)

RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); PRP (Properties); ANST (Analytical study); BIOL (Biological study); USES

(for LMP protein cDNA; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk

injury)

IT cDNA sequences

(for LMP proteins of human and rat; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Drug delivery systems

(implants, LMP protein in; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Drug delivery systems

(injections, LMP protein in; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Spinal column, disease

(intervertebral disk hernia; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Spinal column

(intervertebral disk; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Spinal cord

(lumbar, fusion, LMP protein in gene therapy in; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Bone formation

(mineralization, LMP protein in; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Molecular cloning

(of LIM mineralization proteins; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Protein sequences

(of LMP proteins of human and rat; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Cell differentiation

(osteoblast, LMP protein in; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Stem cell

(pluripotent, LIM mineralization protein mRNA in; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Cytomegalovirus

(promoter for LMP proteins synthesis; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Mutation

(splice site, LMP protein; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Glycosaminoglycans, biological studies

RL: BSU (Biological study, unclassified); BIOL (Biological study)
(sulfated, LMP protein in inducing synthesis of; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Polymers, biological studies

RL: DEV (Device component use); THU (Therapeutic use); BIOL (Biological

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study); USES (Uses)
        (synthetic, as carrier for LMP protein containing cell used in
        intervertebral disk implant; cDNAs encoding rat and human LIM
        mineralization proteins and their use in treatment of disk degeneration
        and disk injury)
     Spinal column
IT
        (vertebra, annulus fibrosus, nucleus pulposus, LIM mineralization
        protein mRNA in; cDNAs encoding rat and human LIM mineralization
        proteins and their use in treatment of disk degeneration and disk
        injury)
                                 530167-62-5
                                                 530167-64-7
IT
     530167-58-9
                    530167-59-0
     RL: BSU (Biological study, unclassified); PRP (Properties); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (amino acid sequence; cDNAs encoding rat and human LIM mineralization
        proteins and their use in treatment of disk degeneration and disk
        injury)
IT
     256606-43-6, GenBank AC023788
     RL: BSU (Biological study, unclassified); PRP (Properties); BIOL
     (Biological study)
        (cDNAs encoding rat and human LIM mineralization proteins and their use
        in treatment of disk degeneration and disk injury)
                  530167-60-3
                                 530167-61-4 530167-63-6
IT
     530167-57-8
     RL: BSU (Biological study, unclassified); PRP (Properties); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (nucleotide sequence; cDNAs encoding rat and human LIM mineralization
        proteins and their use in treatment of disk degeneration and disk
        injury)
     530167-30-7
IT
                    530167-31-8
     RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); PRP
     (Properties); ANST (Analytical study); BIOL (Biological study); USES
     (Uses)
        (primer sequence; cDNAs encoding rat and human LIM mineralization
        proteins and their use in treatment of disk degeneration and disk
     530170-01-5, 3: PN: WO03042368 SEQID: 3 unclaimed DNA
IT
                   530170-04-8, 6: PN: WO03042368 SEQID: 6 unclaimed DNA
     530170-03-7
     530170-05-9, 7: PN: WO03042368 SEQID: 7 unclaimed DNA 530170-06-0, 8:
     PN: WO03042368 SEQID: 8 unclaimed DNA 530170-07-1 530170-08-2
     530170-09-3 530170-10-6 530170-11-7

530170-14-0 530170-15-1 530170-16-2

530170-19-5 530170-20-8 530170-21-9

530170-24-2 530170-25-3 530170-26-4

530170-30-0 530170-31-1 530170-33-3
                                                 530170-12-8
                                                                530170-13-9
                                                 530170-17-3
                                                                530170-18-4
                                                 530170-22-0
                                                                530170-23-1
                                                 530170-28-6
                                                                530170-29-7
                                                 530170-34-4
                                                                530170-35-5
                   530170-37-7 530170-38-8
                                                 530170-39-9
                                                                530170-40-2
     530170-36-6
     RL: PRP (Properties)
        (unclaimed nucleotide sequence; cDNAs encoding rat and human LIM
        mineralization proteins and their use in treatment of disk degeneration
        and disk injury)
IT
     530170-27-5
     RL: PRP (Properties)
        (unclaimed protein sequence; cDNAs encoding rat and human LIM
        mineralization proteins and their use in treatment of disk degeneration
        and disk injury)
IT
     530158-96-4
     RL: PRP (Properties)
        (unclaimed sequence; cDNAs encoding rat and human LIM mineralization
        proteins and their use in treatment of disk degeneration and disk
        injury)
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L9 ANSWER 4 OF 7 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 2002:181215 HCAPLUS

DOCUMENT NUMBER: 137:358009

TITLE: Optimization of diamine bridges in glutaraldehyde

treated bioprosthetic aortic wall tissue
Human, Paul; Bezuidenhout, Deon; Torrianni,

Mark; Hendriks, Marc; Zilla, Peter

CORPORATE SOURCE: Cape Heart Centre, Department of Cardiothoracic

Surgery, University of Cape Town, Cape Town, S. Afr.

SOURCE: Biomaterials (2002), 23(10), 2099-2103

CODEN: BIMADU; ISSN: 0142-9612

PUBLISHER: Elsevier Science Ltd.

DOCUMENT TYPE: Journal LANGUAGE: English

Objective: Bioprosthetic calcification can be significantly mitigated by AB both increased concns. of glutaraldehyde (GA) and the introduction of diamine (DA) bridges. The purpose of the present study was to evaluate whether an optimal effect of DA-enhanced fixation can be achieved by titration of dialdehyde and diamine concns. Methods: Porcine aortic roots were fixed at 0.05% GA (under-fixation) or 0.2% GA and 0.7% GA (com. fixation). An interim step of DA treatment (1-Lysine; 0, 25, 50 or 100 mm; 37°C; 2 days) was followed by completion of the GA fixation (37°C; 5 days). Aortic wall coupons (12 mm) were punched out and implanted s.c. into seven-week old Long-Evans rats for 60 days. Calcium content was assessed by atomic absorption spectroscopy and histol. Results: Increasing the 1-Lysine concns. beyond 25 mm was essential to achieve the anti-calcific effect of DA-enhanced fixation. This effect was proportional to the GA concns. applied. Compared to non-enhanced GA fixation (0 mm DA), calcification increased by 17.4% (p=0.2114) in 0.05% fixed tissue but decreased by 32.0% (p<0.0001) and 45.1% (p<0.0002) in 0.2% and 0.7% GA, resp., when the DA concentration was 100 mm. Histol. the extent, but not the pattern of calcification, was affected. Conclusion: The calcium mitigating effect of diamine-treatment as an interim step of glutaraldehyde fixation is proportional to the GA concentration applied. Within

com. 0.7% GA fixation 100 mm DA has the potential to practically halve aortic wall calcification.

CC 63-7 (Pharmaceuticals)

ST aorta glutaraldehyde crosslinking calcification

IT Artery

AUTHOR (S):

(aorta; optimization of diamine bridges in glutaraldehyde treated bioprosthetic aortic wall tissue)

IT Prosthetic materials and Prosthetics

(bioprosthetics; optimization of diamine bridges in glutaraldehyde treated bioprosthetic aortic wall tissue)

IT Amines, formation (nonpreparative)

RL: FMU (Formation, unclassified); FORM (Formation, nonpreparative) (diamines; optimization of diamine bridges in glutaraldehyde treated bioprosthetic aortic wall tissue)

IT Calcification

## Crosslinking

(optimization of diamine bridges in glutaraldehyde treated bioprosthetic aortic wall tissue)

IT Heart

(valve, bioprosthetic; optimization of diamine bridges in glutaraldehyde treated bioprosthetic aortic wall tissue)

IT 56-87-1, L-Lysine, biological studies

RL: BSU (Biological study, unclassified); BIOL (Biological study) (optimization of diamine bridges in glutaraldehyde treated bioprosthetic aortic wall tissue)

IT 111-30-8, Glutaraldehyde

RL: MOA (Modifier or additive use); RCT (Reactant); RACT (Reactant or reagent); USES (Uses)

(optimization of diamine bridges in glutaraldehyde treated

bioprosthetic aortic wall tissue)

REFERENCE COUNT:

THERE ARE 20 CITED REFERENCES AVAILABLE FOR THIS 20 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 5 OF 7 HCAPLUS COPYRIGHT 2004 ACS on STN L9

ACCESSION NUMBER:

2000:553459 HCAPLUS

DOCUMENT NUMBER:

133:155511

TITLE:

Highly-mineralized osteogenic sponge compositions, and

uses thereof

INVENTOR (S):

McKay, William F.

PATENT ASSIGNEE(S):

SDGI Holdings, Inc., USA PCT Int. Appl., 34 pp.

SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

J	KIND DATE				i	APPI	LICAT	ION I	DATE										
-																			
V	WO 2000045871					A1 20000810			ī	WO 2	2000-1	US304	20000204						
		W:	ΑE,	AL,	AM,	AΤ,	AU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	CA,	CH,	CN,	CR,	CU,	
			CZ,	DE,	DK,	DM,	EE,	ES,	FI,	GB,	GD,	GE,	GH,	GM,	HR,	ΗU,	ID,	IL,	
			IN,	IS,	JP,	KE,	KG,	ΚP,	KR,	ΚZ,	LC,	LK,	LR,	LS,	LT,	LU,	LV,	ΜA,	
			MD,	MG,	MK,	MN,	MW,	MX,	NO,	NZ,	PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	
			SK,	SL,	ΤJ,	TM,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VN,	YU,	ZA,	ZW,	AM,	
			ΑZ,	BY,	KG,	ΚZ,	MD,	RU,	TJ,	TM									
		RW:	GH,	GM,	ΚE,	LS,	MW,	SD,	SL,	SZ,	TZ,	UG,	ZW,	ΑT,	ΒE,	CH,	CY,	DE,	
			DK,	ES,	FI,	FR,	GB,	GR,	·IE,	IT,	LU,	MC,	NL,	PT,	SE,	BF,	ВJ,	CF,	
			CG,	CI,	CM,	GA,	GN,	GW,	ML,	MR,	NE,	SN,	TD,	TG					
F	EΡ	1150	726			A1		2001	1107	]	EP 2	2000-9	9059	20000204					
I	EΡ	1150	726			B1		2003	1105										
		R:	ΑT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,	
			ΙE,	SI,	LT,	LV,	FI,	RO											
ز	JP 2002536077							2002	1029		JP 2	2000-9	5969	20000204					
I	TΑ	2533				E		2003	1115	7	AT 2	2000-9	9059	89	20000204				
I	AU 772682							2004	0506	7	AU 2	2000-2	2756	20000204					
E	ΞS	2209	820			Т3		2004	701	J	ES 2	2000-9	9059	89	20000204				
PRIOR]	ITY	APP	LN.	INFO	. :					τ	US 1999-118615P					P 19990204			
										1	WO 2	7-000	JS304	43	Ţ	<b>V</b> 20	0000	204	
									•		-							_	

- Osteogenic sponge compns. having enhanced osteoinductive properties for ΔR use in bone repair are described. The compns. include a quickly resorbable porous carrier, a more slowly resorbed mineral scaffold and an osteogenic factor, preferably a bone morphogenetic protein. The compns. enable increased osteoinductive activity while retaining a reliable scaffold for the formation of new bone at an implant site. Methods for therapeutic use of the compns. are also described.
- IC ICM A61L027-22
  - ICS A61L027-56; A61L027-46; A61K038-18
- 63-7 (Pharmaceuticals) CC
  - Section cross-reference(s): 2
- osteogenic sponge morphogenetic protein bone implant ST
- IT Bone morphogenetic proteins
  - RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); DEV (Device component use); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

(2; highly-mineralized osteogenic sponge compns. for repair of bone)

IT Bone morphogenetic proteins

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); DEV (Device component use); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

(7; highly-mineralized osteogenic sponge compns. for repair of bone)

IT Proteins, specific or class

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); DEV (Device component use); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

(LMP (LIM-mineralization proteins); highly-mineralized osteogenic sponge compns. for repair of **bone**)

IT Ceramics

(biocompatible; highly-mineralized osteogenic sponge compns. for repair of bone)

IT Bone formation

Osteoblast

Osteoclast

Particle size distribution

(highly-mineralized osteogenic sponge compns. for repair of bone)

IT Bone morphogenetic proteins

Collagens, biological studies

Platelet-derived growth factors

Steroids, biological studies

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); DEV (Device component use); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

(highly-mineralized osteogenic sponge compns. for repair of bone)

IT Bone

(implant; highly-mineralized osteogenic sponge compns. for repair of bone)

IT Porosity

(microporosity; highly-mineralized osteogenic sponge compns. for repair of bone)

IT Bone marrow

(osteogenic enhancing factor of; highly-mineralized osteogenic sponge compns. for repair of **bone**)

IT Growth factors, animal

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (osteogenins; highly-mineralized osteogenic sponge compns. for repair of bone)

IT Bone

(particles of; highly-mineralized osteogenic sponge compns. for repair of bone)

IT Surgery

(spinal fusion; highly-mineralized osteogenic sponge compns. for repair of bone)

IT Medical goods

(sponges, osteogenic; highly-mineralized osteogenic sponge compns. for repair of bone)

IT Spinal column

(vertebra, fusion of; highly-mineralized osteogenic sponge compns. for repair of **bone**)

IT Transforming growth factors

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); DEV (Device component use); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

 $(\beta$ -; highly-mineralized osteogenic sponge compns. for repair of bone)

IT Microglobulins

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); DEV (Device component use); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

 $(\hat{\beta}\text{-microglobulins}; \text{ highly-mineralized osteogenic sponge compns.}$  for repair of **bone**)

IT 10103-46-5, Calcium phosphate

RL: DEV (Device component use); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

(biocompatible ceramics; highly-mineralized osteogenic sponge compns. for repair of bone)

IT 61912-98-9, Insulin like growth factor 62031-54-3, Fgf
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); DEV (Device component use); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

(highly-mineralized osteogenic sponge compns. for repair of bone)

REFERENCE COUNT:

9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

## L9 ANSWER 6 OF 7 HCAPLUS COPYRIGHT 2004 ACS on STN ACCESSION NUMBER: 1999:139773 HCAPLUS

DOCUMENT NUMBER:

INVENTOR(S):

130:200953

TITLE:

A method of crosslinking collagen-based

material and bioprosthetic devices produced therefrom Hendriks, Marc; Verhoeven, Michel; Cahalan, Patrick

T.; Torrianni, Mark W.; Fouache, Benedicte;

Cahalan, Linda

PATENT ASSIGNEE(S):

Medtronic, Inc., USA

SOURCE:

Eur. Pat. Appl., 26 pp.

CODEN: EPXXDW

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

F	PATENT NO.				KINI	)	DATE			APPL	ICAT	ION 1	D.	DATE				
-														-				
E	EP. 897942				A1 1999			9990224			998-	3065	1	19980818				
E	ΞP	8979	42			B1		2004	0310									
		R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,
			ΙE,	SI,	LT,	LV,	FI,	RO										
Ü	JS	6166	184			A		2000	1226		US 1	997-	9127	78		1	99708	318
PRIORI												997-						318
												rial						
ā	ami	ne g	roup	and	I co	llage	n c	arbo	xyl e	grou	ıps a	re p	rovi	ded.	The	e me	thods	3
-	com	pris	e blo	ockii	ng at	lea	st	a po	rtio	n of	the	col	lage	n am	ine o	grou	ps w	ith a
b	olo	ckin	g age	ent t	co fo	orm h	loc	ked	amin	e gr	oups	; co	ntac	ting	the	col	lager	ı-based

```
material having the blocked amine groups with a polyfunctional spacer; and_
 activating at least a portion of the collagen carboxyl groups after
 blocking at least a portion of the collagen amine groups, wherein the
(polyfunctional spacer crosslinks the collagen-based material and
 wherein said contacting step may be effected before or after said
 activating step. Bioprosthetic devices made from these
crosslinked collagen-based materials are also provided.
 Crosslinking involving the JEFFAMINE spacers shows the fastest
 rehydration, whereas glutaraldehyde crosslinking tends to be a
 bit slower. The highly hydrophilic crosslinked collagen-derived
 materials promote infiltration and diffusion of tissue fluid through the
 material matrix, providing supply of oxygen, nutritive substances,
 electrolytes and drainage of metabolites. Also, ingrowth of capillary
 blood vessels and cells is promoted, 25 and consequently the healing
 response is improved. In addition, hydrophilicity improves the blood
 compatibility of the material. Collagen samples crosslinked
 according to the method of the invention involving the Jeffamine D230
 spacer had a cell growth inhibition of 25%, while cells with a deviant
 morphol. were not observed
 ICM C08H001-06
 ICS A61L027-00
 63-7 (Pharmaceuticals)
 Section cross-reference(s): 45
 crosslinking collagen bioprosthetic device manuf
 Acylation
    (agents; crosslinking collagen-based material for
   bioprosthetic devices manufacture)
Heart
    (aortic valve; crosslinking collagen-based material for
   bioprosthetic devices manufacture)
 Collagens, biological studies
RL: DEV (Device component use); SPN (Synthetic preparation); THU
 (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
 (Uses)
    (crosslinked; crosslinking collagen-based material
    for bioprosthetic devices manufacture)
Biocompatibility
Calcification
  Crosslinking
 Transplant and Transplantation
    (crosslinking collagen-based material for bioprosthetic
   devices manufacture)
Aldehydes, reactions
Azides
Ketones, reactions
RL: NUU (Other use, unclassified); RCT (Reactant); RACT (Reactant or
 reagent); USES (Uses)
    (crosslinking collagen-based material for bioprosthetic
   devices manufacture)
Collagens, biological studies
RL: RCT (Reactant); THU (Therapeutic use); BIOL (Biological study); RACT
 (Reactant or reagent); USES (Uses)
    (crosslinking collagen-based material for bioprosthetic
   devices manufacture)
7732-18-5, Water, processes
RL: PEP (Physical, engineering or chemical process); PROC (Process)
    (absorption; crosslinking collagen-based material for
   bioprosthetic devices manufacture)
1122-58-3, 4-Dimethylaminopyridine
                                     2592-95-2, N-Hydroxybenzotriazole
6066-82-6, N-Hydroxysuccinimide
                                  39743-84-5
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IC

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RL: NUU (Other use, unclassified); USES (Uses)
        (crosslinking collagen-based material for bioprosthetic
       devices manufacture)
                       111-30-8, Glutaraldehyde
                                                  123-38-6, Propanal,
IT
    66-25-1, Hexanal
                                                          530-62-1,
                123-72-8, Butanal 420-04-2, Cyanamide
    reactions
                               538-75-0, N,N'-Dicyclohexylcarbodiimide
    1,1'-Carbonyldiimidazole
    616-02-4, Citraconic anhydride 693-13-0, N,N'-Diisopropylcarbodiimide
    830-03-5, p-Nitrophenyl acetate 1865-01-6, p-Nitrophenyl formate
                                  2491-17-0
                                               2635-84-9, p-Nitrophenyl
    2466-76-4, 1-Acetylimidazole
    butyrate 6066-82-6D, N-Hydroxysuccinimide, esters 9046-10-0, Jeffamine
            14464-29-0, N-Hydroxysuccinimidyl acetate 16357-59-8,
    2-Ethoxy-1-ethoxycarbonyl-1,2-dihydroquinoline
                                                     25952-53-8,
    1-Ethyl-3-(3-dimethylaminopropyl)carbodiimide hydrochloride
                                                                  30364-55-7
    74124-79-1, N,N'-Disuccinimidyl carbonate 94820-31-2
                                                             152305-87-8
    RL: NUU (Other use, unclassified); RCT (Reactant); RACT (Reactant or
    reagent); USES (Uses)
        (crosslinking collagen-based material for bioprosthetic
       devices manufacture)
    74-94-2, Dimethylamine borane 75-22-9, Trimethylamine borane
TΤ
    16940-66-2, Sodium borohydride 25895-60-7, Sodium cyanoborohydride
    65605-36-9
```

RL: RCT (Reactant); RACT (Reactant or reagent) (crosslinking collagen-based material for bioprosthetic devices manufacture)

REFERENCE COUNT:

THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 7 OF 7 HCAPLUS COPYRIGHT 2004 ACS on STN ACCESSION NUMBER: 1999:7871 HCAPLUS

6

DOCUMENT NUMBER:

130:57274

TITLE:

SOURCE:

Bone graft composites and spacers

McKay, William F. INVENTOR(S):

PATENT ASSIGNEE(S):

SDGI Holdings, Inc., USA

PCT Int. Appl., 56 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PA	PATENT NO.						KIND DATE				ICAT:	DATE							
						-													
WO	WO 9856433					A1 19981217			1	WO 19	998-1		19980611						
	W:	AL,	AM,	AT,	AU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	CA,	CH,	CN,	CU,	CZ,	DE,		
		DK,	EE,	ES,	FI,	GB,	GE,	GH,	GM,	GW,	HU,	ID,	IL,	IS,	JP,	ΚE,	KG,		
		ΚP,	KR,	ΚZ,	LC,	LK,	LR,	LS,	LT,	LU,	LV,	MD,	MG,	MK,	MN,	MW,	MX,		
		NO,	NZ,	PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,	ТJ,	TM,	TR,	TT,		
		UA,	ŪĠ,	US,	UZ,	VN,	ΥU,	ZW,	AM,	ΑZ,	BY,	KG,	ΚZ,	MD,	RU,	TJ,	TM		
	RW:	GH,	GM,	KE,	LS,	MW,	SD,	SZ,	UG,	ZW,	AT,	BE,	CH,	CY,	DE,	DK,	ES,		
		FI,	FR,	GB,	GR,	ΙE,	IT,	LU,	MC,	NL,	PT,	SE,	BF,	ВJ,	CF,	CG,	CI,		
							ΝE,												
AU	AU 9878185								7	AU 19	998-		19980611						
AU	7382	18			B2	:	2001	0913											
EP	9880	70			A1	:	2000	0329	1	EP 19	998-9		19980611						
EP	9880	70			B1 20040915														
	R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,		
		ΙE,	FΙ																
JP	JP 2002503992						2002	0205	j	JP 19	999-		19980611						
US	6261	586			B1	:	2001	0717	τ	JS 19	999-3		19990831						
PRIORITY	Y APP	LN.	INFO	. :					τ	US 1997-873276						A 19970611			

WO 1998-US11611 W 19980611

- AB A bone graft substitute including a composition of natural selectively deactivated bone material which has been processed to remove associated non-collagenous bone proteins, said bone material containing native collagen materials and naturally associated bone minerals and substantially free from native non-collagenous protein, and a therapeutically effective amount to stimulate bone growth of a bone growth factor in synergistic combination with said bone material. Spacers composed of the bone graft substitute composition and methods for using the spacers are also provided. A diaphysial cortical bone dowel was prepared as well as deactivated allograft and its composite with BMP-2 composite.
- IC ICM A61L027-00
- CC 63-7 (Pharmaceuticals)
- ST bone graft composite spacer
- IT Bone morphogenetic proteins

RL: PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

(2; bone graft composites and spacers)

IT Bone

(artificial; bone graft composites and spacers)

IT Collagens, biological studies

Proteins, general, biological studies

RL: BOC (Biological occurrence); BSU (Biological study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); OCCU (Occurrence); PROC (Process); USES (Uses)

(bone graft composites and spacers)

IT Growth factors, animal

RL: BOC (Biological occurrence); BSU (Biological study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); OCCU (Occurrence); PROC (Process); USES (Uses)

(bone-derived; bone graft composites and spacers)

IT Transplant and Transplantation

Transplant and Transplantation

(bone; bone graft composites and spacers)

IT Prosthetic materials and Prosthetics

(composites, implants; bone graft composites and spacers)

IT Bone

Bone

(transplant; bone graft composites and spacers)

IT 1306-06-5, Hydroxyapatite 7758-87-4, Tricalcium phosphate RL: PEP (Physical, engineering or chemical process); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

(bone graft composites and spacers)

REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT